

**CLAIMS:**

1. An isolated nucleic acid molecule comprising a nucleic acid sequence encoding a polypeptide selected from the group consisting of the lectin domain of a mammalian polypeptide GalNAc-transferase, and a lectin-functional variant or fragment of said lectin domain, wherein said polypeptide does not encompass the intact, functioning catalytic domain of the enzyme.
2. A nucleic acid molecule according to claim 1 comprising a nucleic acid sequence selected from the group consisting of the nucleic acid sequences encoding the GalNAc-T1 to -T16 lectin domains set forth in Table III herein and lectin-functional variants and fragments thereof.
3. The nucleic acid of claim 2 further comprising 30-60 nucleotides of the corresponding GalNAc-transferase sequence at its 5' or 3' end.
4. The nucleic acid of claim 1 wherein the polypeptide GalNAc-transferase or lectin-functional variant or fragment of said lectin domain is human.
5. An isolated lectin polypeptide comprising the lectin domain of a mammalian polypeptide GalNAc-transferase or a lectin-functional variant or fragment thereof.
6. A lectin polypeptide according to claim 5 having an amino acid sequence selected from the group consisting of the amino acid sequences of GalNAc-T1 to -T16 set forth in Table III herein and lectin-functional variants and fragments thereof.
7. The polypeptide of claim 6 further comprising 10-20 amino acid residues of the the corresponding GalNAc-transferase sequence at its carboxy or amino terminus.
8. The polypeptide of claim 5 wherein the polypeptide GalNAc-transferase or a lectin-functional variant or fragment thereof is human.





21. An inhibitor according to claim 18 wherein R is selected from the group consisting of benzyl, phenyl, p-nitrophenyl, umbelliferyl, and naphthalenemethanol.

22. A method of inhibiting mucin secretion in a subject comprising administering an effective amount of a compound that binds to one or more lectin domains  
5 of members of a mammalian family of polypeptide GalNAc-transferases and inhibit binding of such domains to carbohydrates.

23. A method of inhibiting hypersecretion and accumulation of mucin in the lungs of a mammal suffering from a chronic obstructive respiratory pulmonary disease comprising administering to said mammal an effective amount of at least one agent that  
10 inhibits the binding of polypeptide GalNAc-transferase lectin domains to GalNAc-glycopeptides, wherein said agent is selected from the group consisting of GalNAc $\beta$ 1-benzyl, a carbohydrate portion of GalNAc $\beta$ 1-benzyl, a glycoconjugate that includes a carbohydrate portion of GalNAc $\beta$ 1-benzyl or a derivative of either that inhibits the binding of GalNAc-glycopeptides to a GalNAc-transferase lectin domain.

15 24. The method of claim 23 wherein the agent is a glycoconjugate that includes a carbohydrate portion of GalNAc $\beta$ 1-benzyl.

25. The method of claim 23 wherein said mammal is a human.

26. A method of inhibiting the secretion of mucin in a patient comprising administering to the patient a therapeutically effective amount of an agent selected from the  
20 group consisting of GalNAc $\beta$ 1-benzyl, a carbohydrate portion of GalNAc $\beta$ 1-benzyl, a glycoconjugate that includes a carbohydrate portion of GalNAc $\beta$ 1-benzyl or a derivative of either that inhibits the binding of GalNAc-glycopeptides to a GalNAc-transferase lectin domain.

27. The method of claim 26, which selectively inhibits one or more members of  
25 the GalNAc-transferase family without inhibiting other glycosyltransferases selected from

the group consisting of core 1  $\beta$ 1,3-galactosyltransferases,  $\alpha$ 2,6-sialyltransferases, and glycosyltransferases functioning in the O-glycosylation pathway.

28. The method of claim 26 wherein the patient has a disease selected from the group consisting of chronic obstructive pulmonary diseases, asthma, and cystic fibrosis.

5 29. A method of modulating the function of one or more lectin domains of a polypeptide GalNAc-transferase comprising administering an effective amount of GalNAc $\beta$ 1-R which is effective in modulating functions mediated by said lectin domains.

30. The method of claim 29 wherein R represents an aglycone.

31. The method of claim 29 wherein R represents an aryl group.

10 32. The method of claim 30 wherein R is selected from the group consisting of benzyl, phenyl, p-nitrophenyl, umbelliferyl, and naphthalenemethanol.

33. A method of screening one or more test substances for the ability to inhibit or modulate intracellular transport and/or cell surface expression of mucins, O-glycosylated glycoproteins, glycoproteins and proteins in a cell-based assay, which comprises:

15 (i) contacting a cell that expresses mucins, O-glycosylated glycoproteins, glycoproteins and proteins, with one or more test substances under assay conditions suitable for the detection of inhibition or modulation of said expression; and

(ii) measuring whether intracellular transport and cell surface expression of said mucins, O-glycosylated glycoproteins, glycoproteins and proteins are thereby inhibited or  
20 modulated by one or more of the substances.

34. A method of screening one or more test substances for the ability to inhibit or modulate secretions of mucins, O-glycosylated glycoproteins, glycoproteins and proteins in a cell-based assay, which comprises:

(i) contacting a cell that secretes mucins, O-glycosylated glycoproteins, glycoproteins with one or more test substances under assay conditions suitable for the detection of inhibition or modulation of said secretion; and

(ii) measuring whether secretion of said mucins, O-glycosylated glycoproteins, glycoproteins and proteins are thereby inhibited or modulated by one or more of the substances.

35. The method of claim 22, wherein the compound is GalNAc $\beta$ 1-benzyl.

36. The method of claim 23, wherein the compound is GalNAc $\beta$ 1-benzyl.

37. The method of claim 23, wherein the compound is GalNAc $\beta$ 1-benzyl.

38. The method of claim 34, wherein step (ii) further comprises measuring whether the intracellular accumulation of said mucins, O-glycosylated glycoproteins and proteins is inhibited or modulated.

39. A method of inhibiting mucin secretion in a cell comprising delivering to a cell an effective amount of a compound that binds to one or more lectin domains of members of a mammalian family of polypeptide GalNAc-transferases and inhibit binding of such domains to carbohydrates.

40. The method of claim 39, wherein the compound is GalNAc $\beta$ 1-benzyl.